

**Claims:**

1. A method for the non-invasive early detection of colon cancer and/or intestinal cancer precursor cells by means of mutational analysis of the genes for APC, K-ras,  $\beta$ -catenin and B-raf in a sample, characterized in that the method comprises the following steps:
    - collecting a stool and/or tissue sample,
    - homogenizing the sample,
    - obtaining DNA from the sample,
    - performing an amplification reaction in the genes for APC, K-ras,  $\beta$ -catenin and B-raf, using the primers
      - s1        TTGCAGTTATGGTCAATACCC
      - as1       GTGCTCTCAGTATAAACAGGATAAG
      - s2        CCTCAAAGGCTGCCACTTG
      - as2       CTGTGACACTGCTGGAACCTTCGC
      - s3        AGCACCTAGAACCAAATCCAGCAG
      - as3       TGGCATGGTTTGTCCAGGGC
      - s4        ACAAACCATGCCACCAAGCAGA
      - as4       GAGCACTCAGGCTGGATGAACAAG
      - s5        TTCCAGATGCTGATACTTTA
      - as5       CTGAATCATCTAATAGGTCC
- for APC, the primers
- s        CTGGTGGAGTATTTGATAGTG
  - as       TCTATTGTTGGATCATATTC
- for K-ras, the primers
- s        CTGATTTGATGGAGTTGGAC
  - as       CTTGAGTGAAGGACTGAGA
- for  $\beta$ -catenin, and the primers
- s        TGTATCACCATCTCCATATC
  - as       GCATTCTGATGACTTCTGGT
- for B-raf,
- wherein amplification products are formed, and

- performing a mutational analysis in the amplification products.
2. The method according to claim 1, characterized in that the detection of mutations in selected sections of the genes for APC, K-ras,  $\beta$ -catenin and B-raf is effected by means of a DNA chip, said DNA chip including probes for APC, K-ras,  $\beta$ -catenin and B-raf from those regions of the above-mentioned genes that are flanked by the primer sequences specified in claim 1.
  3. The method according to claim 1 or 2, characterized in that the APC, K-ras,  $\beta$ -catenin and B-raf genes are accumulated from total DNA by hybridizing sequence-specific biotinylated oligonucleotides with the genes for APC, K-ras,  $\beta$ -catenin and B-raf using coupling of the biotin residue to streptavidin and subsequent separation via magnetic particles.
  4. The method according to claims 1 to 3, characterized in that amplification products, especially PCR products, are separated in an agarose gel for control purposes prior to purification.
  5. The method according to any of claims 1 to 4, characterized in that the mutational analysis of the PCR products is effected using electrophoretic techniques, preferably SSCP, alternatively by means of a chromatographic procedure, preferably an HPLC-based procedure.

6. The method according to the preceding claim, characterized in that detected mutagenic conformations of a single strand are isolated and optionally sequenced.
7. Primer sequences selected from the group comprising:  
the primers  
s1        TTGCAGTTATGGTCAATACCC  
as1       GTGCTCTCAGTATAAACAGGATAAG  
s2        CCTCAAAGGCTGCCACTTG  
as2       CTGTGACACTGCTGGAACCTTCG  
s3        AGCACCTAGAACCAAATCCAGCAG  
as3       TGGCATGGTTTGTCCAGGGC  
s4        ACAAACCATGCCACCAAGCAGA  
as4       GAGCACTCAGGCTGGATGAACAAG  
s5        TTCCAGATGCTGATACTTTA  
as5       CTGAATCATCTAATAGGTCC  
or alternatively  
s2        GAATCAGCTCCATCCAAGT  
as2       TTTCTGCTATTTGCAGGGT  
for APC, the primers  
s        CTGGTGGAGTATTTGATAGTG  
as       TCTATTGTTGGATCATATTCG  
for K-ras, the primers  
s        CTGATTTGATGGAGTTGGAC  
as       CTTGAGTGAAGGACTGAGAA  
for  $\beta$ -catenin, and the primers  
s        TGTATCACCATCTCCATATC  
as       GCATTCTGATGACTTCTGGT  
for B-raf.
8. Use of the primer sequences according to claim 7 in mutational analysis, especially in the analysis of the APC, K-ras,  $\beta$ -catenin and B-raf genes.

9. A kit, comprising primers selected from the group comprising:

the primers

s1        TTGCAGTTATGGTCAATACCC  
as1        GTGCTCTCAGTATAAACAGGATAAG  
s2        CCTCAAAGGCTGCCACTTG  
as2        CTGTGACACTGCTGGAACCTTCGC  
s3        AGCACCTAGAACCAATCCAGCAG  
as3        TGGCATGGTTTGTCCAGGGC  
s4        ACAAACCATGCCACCAAGCAGA  
as4        GAGCACTCAGGCTGGATGAACAAG  
s5        TTCCAGATGCTGATACTTTA  
as5        CTGAATCATCTAATAGGTCC

or alternatively

s2        GAATCAGCTCCATCCAAGT  
as2        TTTCTGCTATTTGCAGGGT

for APC, the primers

s        CTGGTGGAGTATTTGATAGTG  
as        TCTATTGTTGGATCATATTCG

for K-ras, the primers

s        CTGATTTGATGGAGTTGGAC  
as        CTTGAGTGAAGGACTGAGAA

for  $\beta$ -catenin, and the primers

s        TGTATCACCATCTCCATATC  
as        GCATTCTGATGACTTCTGGT

for B-raf,

and optionally information relating to combining the contents of the kit.

10. Use of the kit according to claim 9 in the detection of colon cancer and/or colon cancer precursor cells.